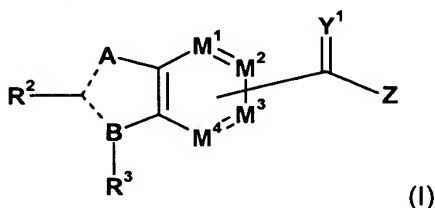


CLAIMS

1. An isomer, enantiomer, diastereoisomer, or tautomer of a compound, represented by formula I:



wherein

----- represents either a single or a double bond;

B is -N- and **A** is =CR¹- or =N-; or

B is =C- and **A** is O, S or NR¹;

- 15 **R**¹ is selected from the group consisting of: H, (C₁₋₆)alkyl optionally substituted with:
 halogen, OR¹¹, SR¹¹ or N(R¹²)₂, wherein **R**¹¹ and each **R**¹² is independently
 H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₆)alkyl-(C₃₋₇)cycloalkyl, (C₁₋₆)alkyl-aryl or
 (C₁₋₆)alkyl-Het, said aryl or Het optionally substituted with **R**¹⁶⁰; or
 20 both **R**¹² are covalently bonded together and to the nitrogen to which they are
 both attached to form a 5, 6 or 7-membered saturated heterocycle;

the group -C(=Y¹)-Z is covalently linked to either **M**² or **M**³,

- 25 **M**¹ is CR^{4a},
M² or **M**³, when not linked to -C(=Y¹)-Z, is CR⁵,
M⁴ is CR^{4b},

- and in addition one or two of the groups selected from **M**¹, **M**², **M**³ and **M**⁴ may also
 30 be N, with the proviso that the group **M**² or **M**³ to which -C(=Y¹)-Z is linked is a C-

atom,

Y^1 is O or S;

- 5 **Z** is defined as $NR^{N2}-SO_2-R^C$ or $NR^{N3}-SO_2-N(R^{N2})R^{N1}$, wherein R^C , R^{N1} or any heterocycle formed by R^{N1} and R^{N2} is optionally substituted with R^{60} ;

R^2 is selected from: halogen or R^{21} , wherein R^{21} is aryl or Het, said R^{21} is optionally substituted with R^{150} ;

10

R^3 is selected from (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl, (C₅₋₇)cycloalkenyl, (C₁₋₃)alkyl-(C₅₋₇)cycloalkenyl, (C₆₋₁₀)bicycloalkyl, (C₁₋₃)alkyl-(C₆₋₁₀)bicycloalkyl, (C₆₋₁₀)bicycloalkenyl, (C₁₋₃)alkyl-(C₆₋₁₀)bicycloalkenyl, **HCy** or (C₁₋₃)alkyl-**HCy**,

15

wherein **HCy** is a saturated or unsaturated 4 to 7-membered heterocyclic group with 1 to 3 heteroatoms selected from O, S and N; said alkyl, cycloalkyl, cycloalkenyl, bicycloalkyl, bicycloalkenyl, **HCy** and alkyl-**HCy** being optionally substituted with from 1 to 4 substituents selected from: a) halogen;

20

b) (C₁₋₆)alkyl optionally substituted with:

- 1 to 3 substituents selected from halogen;
- OR^{31} or SR^{31} wherein R^{31} is H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; or
- $N(R^{32})_2$ wherein each R^{32} is independently H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; or both R^{32} are covalently bonded together and to the nitrogen to which they are attached to form a 5, 6 or 7-membered saturated heterocycle;

25

c) OR^{33} or SR^{33} wherein R^{33} is H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl;

30

d) $N(R^{35})_2$ wherein each R^{35} is independently H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; or both R^{35} are covalently bonded together and to the nitrogen to which they are attached to form a 5, 6 or 7-membered saturated heterocycle;

R^{4a} , R^{4b} , R^5 each are independently H or defined as R^{150} ;

R^{60} is defined as 1 to 4 substituents independently selected from:

- 5 - 1 to 3 substituents selected from halogen;
- one of each substituent selected from: OPO_3H , NO_2 , cyano, azido,
 $C(=NH)NH_2$, $C(=NH)NH(C_{1-6})alkyl$ or $C(=NH)NHCO(C_{1-6})alkyl$, SO_3H ; and
- 1 to 3 substituents selected from:
 - 10 a) $(C_{1-6})alkyl$, $(C_{3-7})cycloalkyl$, $(C_{3-7})spirocycloalkyl$ optionally containing 1 or
 2 heteroatoms selected from N, O and S; $(C_{2-6})alkenyl$, $(C_{2-8})alkynyl$,
 $(C_{1-6})alkyl-(C_{3-7})cycloalkyl$, all of which optionally being substituted with
 R^{150} ;
 - b) OR^O ;
 - c) $OC(O)R^O$;
 - 15 d) SR^O , SO_2R^C , $SO_2N(R^{N2})R^{N1}$, $SO_2N(R^{N2})C(O)R^C$, $CONR^{N3}SO_2N(R^{N2})R^{N1}$,
 or $CONR^{N2}SO_2R^C$;
 - e) $N(R^{N2})R^{N1}$, $N(R^{N2})COOR^C$, $N(R^{N2})SO_2R^C$ or $N(R^{N1})OR^O$;
 - f) $N(R^{N2})COR^C$;
 - g) $N(R^{N3})CON(R^{N2})R^{N1}$;
 - 20 h) $N(R^{N3})COCOR^C$, $N(R^{N3})COCOOR^O$, $N(R^{N3})COCON(R^{N2})OR^O$, or
 $N(R^{N3})COCON(R^{N2})R^{N1}$;
 - i) COR^O ;
 - j) $COOR^O$;
 - k) $CON(R^{N2})R^{N1}$;
 - 25 l) aryl, Het, $(C_{1-4})alkyl-aryl$ or $(C_{1-4})alkyl-Het$, all of which optionally being
 substituted with R^{150} ;

wherein said R^{N1} , R^C and/or R^O are optionally substituted with R^{150} as defined,

30 R^{150} is defined as 1 to 4 substituents independently selected from:

- 1 to 3 substituents selected from halogen;
- one of each substituent selected from: OPO_3H , NO_2 , cyano, azido, SO_3H
 $C(=NH)NH_2$, $C(=NH)NH(C_{1-6})alkyl$ or $C(=NH)NHCO(C_{1-6})alkyl$; and
- 1 to 3 substituents selected from:

- a) (C₁₋₆) alkyl, (C₃₋₇)cycloalkyl, (C₃₋₇)spirocycloalkyl optionally containing 1 or 2 heteroatoms selected from N, O and S; (C₂₋₆)alkenyl, (C₂₋₈)alkynyl, (C₁₋₃) alkyl-(C₃₋₇)cycloalkyl, all of which optionally substituted with R¹⁶⁰;
- b) OR⁰;
- 5 c) OC(O)R⁰;
- d) SR⁰, SO₂R^C, SO₂N(R^{N2})R^{N1} or SO₂N(R^{N2})C(O)R^C;
- e) N(R^{N2})R^{N1}, N(R^{N2})COOR^C, N(R^{N2})SO₂R^C or N(R^{N1})OR⁰;
- f) N(R^{N2})COR^C;
- g) N(R^{N3})CON(R^{N2})R^{N1};
- 10 h) N(R^{N3})COCOR^C, N(R^{N3})COCOOR⁰, N(R^{N3})COCON(R^{N2})OH, N(R^{N3})COCON(R^{N2})O(C₁₋₄)alkyl or N(R^{N3})COCON(R^{N2})R^{N1};
- i) COR⁰;
- j) COOR⁰;
- k) tetrazole, triazole, CONR^{N2}SO₂R^C, CONR^{N3}-SO₂N(R^{N2})R^{N1} or
- 15 CON(R^{N2})R^{N1};
- wherein said R^{N1}, R^C and/or R⁰ are optionally substituted with R¹⁶⁰ as defined;

R¹⁶⁰ is defined as 1, 2 or 3 substituents independently selected from:

20 - 1, 2 or 3 fluorine substituents; and

- one of each substituent selected from tetrazole, triazole, chlorine, bromine, iodine, CN, nitro, (C₁₋₄)alkyl, OCF₃, SCF₃, CF₃, COOR¹⁶¹, SO₃H, SR¹⁶¹, SO₂R¹⁶³, OR¹⁶¹, N(R¹⁶²)₂, SO₂N(R¹⁶²)₂, SO₂NR¹⁶²COR¹⁶², NR¹⁶²SO₂R¹⁶³, -NR¹⁶¹-CO-COOR¹⁶¹, -NR¹⁶¹-CO-CO(NR¹⁶²)₂, -CONR¹⁶¹SO₂R^C, CONR¹⁶¹-

25 SO₂N(R¹⁶²)₂ or -SO₂-NR¹⁶¹-COR^C, NR¹⁶²COR¹⁶² or CON(R¹⁶²)₂, wherein R¹⁶¹, R¹⁶³ and each R¹⁶² is independently (C₁₋₄)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; and R¹⁶¹ and each R¹⁶² may each independently also be H; or both R¹⁶² are covalently bonded together and to the nitrogen to which they are attached to form a 5, 6 or 7-membered saturated heterocycle;

30 R⁰, R^C are independently defined as (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₄)alkyl-(C₃₋₇)cycloalkyl, (C₂₋₆)alkenyl, aryl, Het, (C₁₋₄)alkyl-aryl, or (C₁₋₄)alkyl-Het; or R⁰ is also optionally defined as H.

R^{N1} is H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₄)alkyl-(C₃₋₇)cycloalkyl, (C₂₋₆)alkenyl, aryl, Het, (C₁₋₄)alkyl-aryl, (C₁₋₄)alkyl-Het; and

5 R^{N2} , R^{N3} , R^{N4} are independently H, CH₃, (C₂₋₆)alkyl, (C₃₋₆)cycloalkyl, (C₁₋₄)alkyl-(C₃₋₆)cycloalkyl; all of which being optionally substituted with halogen, carboxy or (C₁₋₆)alkoxycarbonyl; and/or wherein said alkyl, cycloalkyl or alkylcycloalkyl is optionally substituted with hydroxy, (C₁₋₆)alkyl, (C₁₋₆)alkoxy, amino, -NH(C₁₋₄)alkyl and/or -N((C₁₋₄)alkyl)₂; or

10 in the case

a) of a group N(R^{N2}) R^{N1} the substituents R^{N2} and R^{N1} ; or
b) of a group NR^{N3}-N(R^{N2}) R^{N1} the substituents R^{N3} and R^{N1} , or R^{N2} and R^{N1} ; may be covalently bonded together to form a 4-, 5-, 6- or 7-membered saturated or unsaturated N-containing heterocycle or a 8-, 9-, 10- or 11-membered N-containing heterobicycle, each optionally having additionally
15 from 1 to 3 heteroatoms selected from O, N, and S;

wherein Het is defined as a 4-, 5-, 6- or 7-membered heterocycle having 1 to 4 heteroatoms selected from O, N and S, or a 8-, 9-, 10- or 11-membered heterobicycle having 1 to 5 heteroatoms selected from O, N and S;
20

or a salt thereof.

2. The compound according to claim 1, wherein

25

----- represents either a single or a double bond;

B is -N- and A is CR¹ or =N-; or

30 B is =C- and A is O, S or NR¹;

R^1 is selected from the group consisting of: H, (C₁₋₆)alkyl optionally substituted with:
halogen, OR¹¹, SR¹¹ or N(R^{12})₂, wherein R^{11} and each R^{12} is independently
35 H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₆)alkyl-(C₃₋₇)cycloalkyl, (C₁₋₆)alkyl-aryl or

(C₁₋₆)alkyl-Het, said aryl or Het optionally substituted with R¹⁶⁰; or
both R¹² are covalently bonded together and to the nitrogen to which they are
both attached to form a 5, 6 or 7-membered saturated heterocycle;

5 the group -C(=Y¹)-Z is covalently linked to either M² or M³,

M¹ is CR^{4a},

one of M² and M³ is CR⁵,

M⁴ is CR^{4b},

10

and in addition one or two of the groups selected from M¹, M², M³ and M⁴ may also
be N, with the proviso that the group M² or M³ to which -C(=Y¹)-Z is linked is an C-
atom,

15 Y¹ is O or S;

Z is defined as NR^{N2}-SO₂-R^C, wherein R^C is optionally substituted with R⁶⁰;

20 R² is selected from: halogen or R²¹, wherein R²¹ is aryl or Het, said R²¹ is optionally
substituted with R¹⁵⁰;

R³ is selected from (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl,
(C₅₋₇)cycloalkenyl, (C₁₋₃)alkyl-(C₅₋₇)cycloalkenyl, (C₆₋₁₀)bicycloalkyl, (C₁₋₃)alkyl-
(C₆₋₁₀)bicycloalkyl, (C₆₋₁₀)bicycloalkenyl, (C₁₋₃)alkyl-(C₆₋₁₀)bicycloalkenyl, HCy
25 or (C₁₋₃)alkyl-HCy,

wherein HCy is a saturated or unsaturated 4 to 7-membered heterocyclic
group with 1 to 3 heteroatoms selected from O, S and N;

said alkyl, cycloalkyl, cycloalkenyl, bicycloalkyl, bicycloalkenyl, HCy and
alkyl-HCy being optionally substituted with from 1 to 4 substituents selected
30 from: a) halogen;

b) (C₁₋₆)alkyl optionally substituted with:

- OR³¹ or SR³¹ wherein R³¹ is H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or
(C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; or
- N(R³²)₂ wherein each R³² is independently H, (C₁₋₆)alkyl,

(C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; or both R³² are covalently bonded together and to the nitrogen to which they are attached to form a 5, 6 or 7-membered saturated heterocycle;

- 5 c) OR³³ or SR³³ wherein R³³ is H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl;
- d) N(R³⁵)₂ wherein each R³⁵ is independently H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl or (C₁₋₃)alkyl-(C₃₋₇)cycloalkyl; or both R³⁵ are covalently bonded together and to the nitrogen to which they are attached to
- 10 form a 5, 6 or 7-membered saturated heterocycle;

R^{4a}, R^{4b}, R⁵ each are independently H or defined as R¹⁵⁰;

R⁶⁰ is defined as 1 to 4 substituents independently selected from:

- 15 - 1 to 3 substituents selected from halogen;
- one of each substituent selected from: OPO₃H, NO₂, cyano, azido, C(=NH)NH₂, C(=NH)NH(C₁₋₆)alkyl or C(=NH)NHCO(C₁₋₆)alkyl, SO₃H; and
- 1 to 3 substituents selected from:
- 20 a) (C₁₋₆) alkyl, (C₃₋₇)cycloalkyl, C₃₋₇ spirocycloalkyl optionally containing 1 or 2 heteroatom selected from N, O and S; (C₂₋₆)alkenyl, (C₂₋₈)alkynyl, (C₁₋₆)alkyl-(C₃₋₇)cycloalkyl, all of which optionally being substituted with R¹⁵⁰;
- b) OR⁰;
- c) OC(O)R⁰;
- 25 d) SR⁰, SO₂R^C, SO₂N(R^{N2})R^{N1}, SO₂N(R^{N2})C(O)R^C or CONR^{N2}SO₂R^C;
- e) N(R^{N2})R^{N1}, N(R^{N2})COOR^C or N(R^{N2})SO₂R^C;
- f) N(R^{N2})COR^C;
- g) N(R^{N3})CON(R^{N2})R^{N1};
- h) N(R^{N3})COCOR^C, N(R^{N3})COCOOR⁰ or N(R^{N3})COCON(R^{N2})R^{N1};
- 30 i) COR⁰;
- j) COOR⁰;
- k) CON(R^{N2})R^{N1};
- l) aryl, Het, (C₁₋₄)alkyl)aryl or (C₁₋₄)alkyl)Het, all of which optionally being substituted with R¹⁵⁰;

wherein said R^{N1} , R^C and/or R^O are optionally substituted with R^{150} as defined,

R^{150} is defined as 1 to 4 substituents independently selected from:

- 5 - 1 to 3 substituents selected from halogen;
 - one of each substituent selected from: OPO_3H , NO_2 , cyano, azido, $C(=NH)NH_2$, $C(=NH)NH(C_{1-6})alkyl$ or $C(=NH)NHCO(C_{1-6})alkyl$; and
 - 1 to 3 substituents selected from:
 - 10 a) $(C_{1-6})alkyl$, $(C_{3-7})cycloalkyl$, C_{3-7} spirocycloalkyl optionally containing 1 or 2 heteroatoms selected from N, O and S; $(C_{2-6})alkenyl$, $(C_{2-8})alkynyl$, $(C_{1-3})alkyl-(C_{3-7})cycloalkyl$, all of which optionally substituted with R^{160} ;
 - b) OR^O ;
 - c) $OC(O)R^O$;
 - d) SR^O , SO_2R^C , $SO_2N(R^{N2})R^{N1}$ or $SO_2N(R^{N2})C(O)R^C$;
 - 15 e) $N(R^{N2})R^{N1}$, $N(R^{N2})COOR^C$ or $N(R^{N2})SO_2R^C$;
 - f) $N(R^{N2})COR^C$;
 - g) $N(R^{N3})CON(R^{N2})R^{N1}$;
 - h) $N(R^{N3})COCOR^C$, $N(R^{N3})COCOOR^O$ or $N(R^{N3})COCON(R^{N2})R^{N1}$;
 - wherein R^{N1} is as defined or OH, OAlkyl;
 - 20 i) COR^O ;
 - j) $COOR^O$;
 - k) tetrazole or $CON(R^{N2})R^{N1}$;
- wherein said R^{N1} , R^C and/or R^O are optionally substituted with R^{160} as defined;

25

R^{160} is defined as 1, 2 or 3 substituents independently selected from:

- 1, 2 or 3 fluorine substituents; and
- one of each substituent selected from tetrazole, chlorine, bromine, iodine, CN, nitro, $C_{1-4}alkyl$, CF_3 , $COOR^{161}$, SO_3H , SR^{161} , SO_2R^{163} , OR^{161} , $N(R^{162})_2$, $SO_2N(R^{162})_2$, $SO_2NR^{162}COR^{162}$, $NR^{162}SO_2R^{163}$, $NR^{162}COR^{162}$ or $CON(R^{162})_2$,
 - 30 wherein R^{161} , R^{163} and each R^{162} is independently $(C_{1-4})alkyl$, $(C_{3-7})cycloalkyl$ or $(C_{1-3})alkyl-(C_{3-7})cycloalkyl$; and R^{161} and each R^{162} may each independently also be H; or both R^{162} are covalently bonded together and to the nitrogen to which they are attached to form a 5, 6 or 7-membered

saturated heterocycle;

R^O , R^C are independently defined as (C₁₋₆)alkyl, (C₃₋₆)cycloalkyl, (C₁₋₄)alkyl-
(C₃₋₆)cycloalkyl, (C₂₋₆)alkenyl, aryl, **Het**, (C₁₋₄)alkyl-aryl, (C₁₋₄)alkyl-**Het**;

5

R^{N1} is H, (C₁₋₆)alkyl, (C₃₋₇)cycloalkyl, (C₁₋₄)alkyl-(C₃₋₆)cycloalkyl, (C₂₋₆)alkenyl, aryl,
Het, (C₁₋₄)alkyl-aryl, (C₁₋₄)alkyl-**Het**; or

R^{N2} , R^{N3} , R^{N4} are independently H, CH₃, (C₂₋₆)alkyl, (C₃₋₆)cycloalkyl, (C₁₋₄)alkyl-
10 (C₃₋₆)cycloalkyl; all of which being optionally substituted with halogen,
carboxy or C₁₋₆-alkoxycarbonyl; and/or wherein said alkyl, cycloalkyl or
alkylcycloalkyl is optionally substituted with hydroxy, C₁₋₆-alkyl, C₁₋₆-alkoxy,
amino, -NH(C₁₋₄-alkyl) and/or -N(C₁₋₄-alkyl)₂; and

15

in the case

a) of a group N(R^{N2}) R^{N1} the substituents R^{N2} and R^{N1} ; or

b) of a group NR^{N3}-N(R^{N2}) R^{N1} the substituents R^{N3} and R^{N1} , or R^{N2} and R^{N1} ;
may be covalently bonded together to form a 4-, 5-, 6- or 7-membered
saturated or unsaturated N-containing heterocycle or a 8-, 9-, 10- or 11-

20

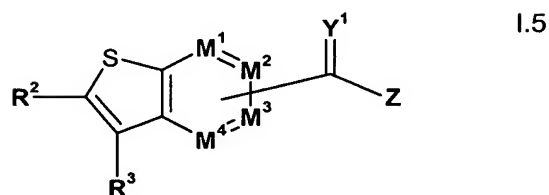
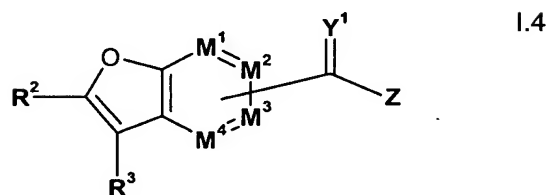
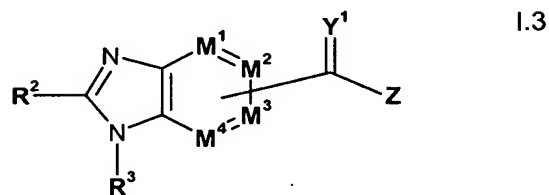
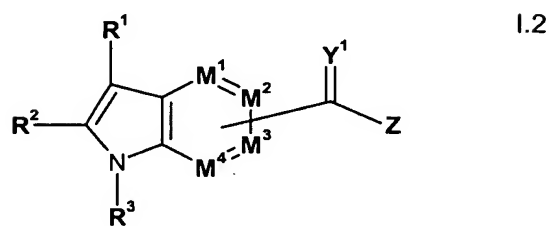
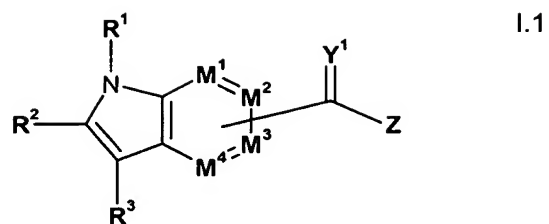
membered N-containing heterobicycle each may have additionally from 1 to 3
heteroatoms selected from O, N, and S, wherein said heterocycle or
heterobicycle is optionally substituted as defined;

wherein **Het** is defined as a 4-, 5-, 6- or 7-membered heterocycle having 1 to 4
25 heteroatoms selected from O, N and S, or a 8-, 9-, 10- or 11-membered
heterobicycle having 1 to 5 heteroatoms selected from O, N and S;

or a salt thereof.

30

3. The compound according to claim 1 selected from the group of formulas I.1
to I.5

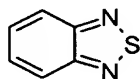


wherein R^1 , R^2 , R^3 , Y^1 , Z , M^1 , M^2 , M^3 and M^4 are defined as in claim 1.

4. The compound according to claim 1, wherein R^1 is selected from the group consisting of: H and (C₁₋₆)alkyl.
5. The compound according to claim 4, wherein R^1 is H, CH₃, ethyl, or isobutyl.

6. The compound according to claim 5, wherein R^1 is H or CH_3 .
7. The compound according to claim 6, wherein R^1 is CH_3 .
- 5 8. The compound according to claim 1, wherein Y^1 is O.
9. The compound according to claim 1, wherein Z is $NR^{N3}-SO_2-N(R^{N2})R^{N1}$, wherein R^{N1} or any heterocycle formed by R^{N1} and R^{N2} is optionally substituted with R^{60} , and wherein R^{N3} , R^{N2} , R^{N1} and R^{60} are defined as in claim 1.
- 10 10. The compound according to claim 1, wherein Z is $NR^{N2}-SO_2-R^C$, wherein R^C is optionally substituted with R^{60} , and wherein Het , R^{N2} , R^C and R^{60} are defined as in claim 1.
- 15 11. The compound according to claim 10, wherein Z is $NH-SO_2-R^C$, wherein R^C is selected from the group consisting of (C_{1-6}) alkyl, (C_{3-6}) cycloalkyl, (C_{1-3}) alkyl- (C_{3-6}) cycloalkyl, (C_{2-6}) alkenyl, phenyl, naphthyl, Het , (C_{1-3}) alkyl-phenyl, (C_{1-3}) alkyl-naphthyl, (C_{1-3}) alkyl- Het , wherein said alkyl, cycloalkyl, alkyl-cycloalkyl, alkenyl, phenyl, naphthyl, Het , alkyl-phenyl, alkyl-naphthyl, or alkyl- Het , are all optionally substituted with 1 to 4 substituents selected from R^{60} , wherein R^{60} and Het are defined as in claim 10.
- 20 12. The compound according to claim 11, wherein Z is $NH-SO_2-R^C$, wherein R^C is selected from the group consisting of methyl, ethyl, n-propyl, i-propyl, cyclopropyl, cyclobutyl, cyclopentyl, cyclopropylmethyl, cyclobutylmethyl, cyclopentylmethyl, pyrrolidine, piperidine, morpholine, thiomorpholine, piperazine, phenyl, naphthyl, benzyl, thiophene, furan, pyrrole, imidazole, pyrazole, oxazole, isoxazole, thiazole, pyridazine, pyrimidine, pyrazine, diazepine, azepine, quinoline, isoquinoline, benzofuran, benzothiophene, benzothiazole, purine, pteridine,
- 25 30

2,1,3-benzothiadiazole

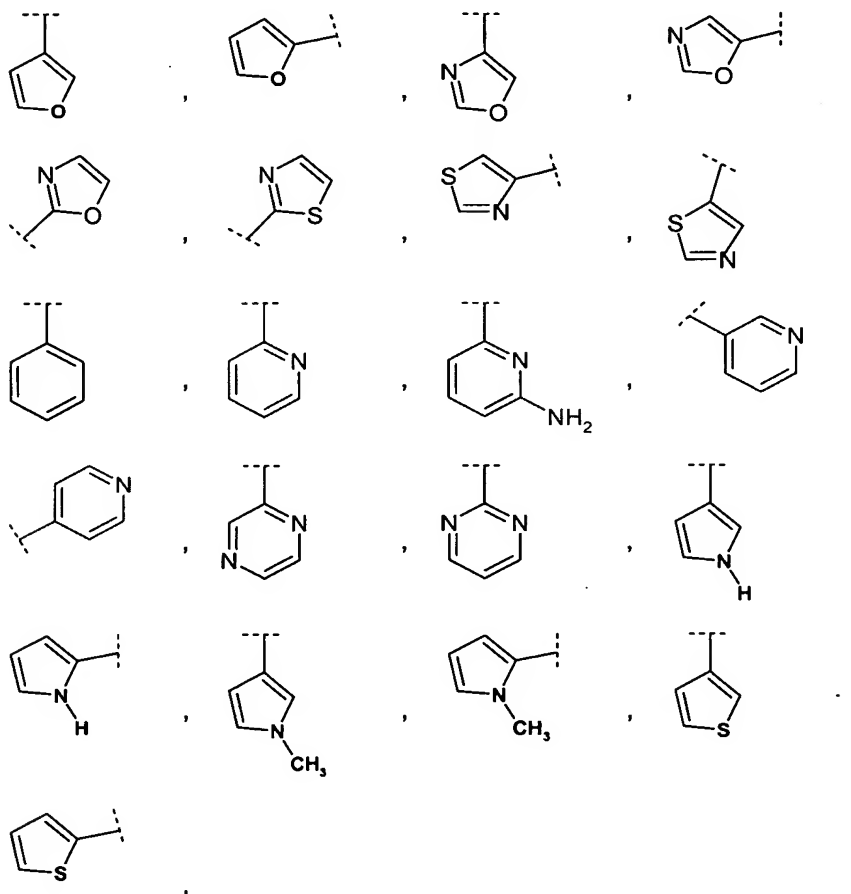


, and



all of which are optionally substituted with 1 to 3 substituents selected from R^{60} , wherein R^{60} is defined as in claim 11.

- 5 **13.** The compound according to claim 1, wherein R^2 is R^{21} , wherein R^{21} is phenyl or Het selected from the group of formulas



and wherein said R^{21} is unsubstituted or substituted with R^{150} , being defined as in claim 1.

10

- 14.** The compound according to claim 1, wherein R^2 is R^{21} , wherein R^{21} is

defined as in claim 1, and wherein R^{21} is optionally substituted with 1, 2 or 3 substituents selected from:

- 1 to 3 substituents selected from halogen;
- one of each substituent selected from: NO_2 , cyano, azido; and
- 1 to 2 substituents selected from:

- a) (C_{1-4}) alkyl or (C_{1-4}) alkoxy, both optionally substituted with OH, $O(C_{1-4})$ alkyl, $SO_2(C_{1-4})$ alkyl, 1 to 3 halogen atoms, amino, $NH(C_{1-4})$ alkyl or $N((C_{1-4})alkyl)_2$;
- b) $NR^{111}R^{112}$ wherein both R^{111} and R^{112} are independently H, (C_{1-4}) alkyl, or R^{112} is (C_{3-7}) cycloalkyl, (C_{1-3}) alkyl (C_{3-7}) cycloalkyl, phenyl, benzyl; or both R^{111} and R^{112} are covalently bonded together and to the nitrogen to which they are attached to form a nitrogen-containing heterocycle, each of said alkyl, cycloalkyl, alkylcycloalkyl, phenyl and benzyl, being optionally substituted with halogen or:
 - OR^{2h} or $N(R^{2h})_2$, wherein each R^{2h} is independently H, (C_{1-4}) alkyl, or both R^{2h} are covalently bonded together and to the nitrogen to which they are attached to form a nitrogen-containing heterocycle;
- c) $NHCOR^{117}$ wherein R^{117} is (C_{1-4}) alkyl, $O(C_{1-4})$ alkyl or $O(C_{3-7})$ cycloalkyl;
- d) and
- e) $CONH_2$, $CONH(C_{1-4})$ alkyl, $CON((C_{1-4})alkyl)_2$.

15. The compound according to claim 1, wherein R^3 is selected from (C_{3-7}) cycloalkyl, (C_{5-7}) cycloalkenyl, (C_{6-10}) bicycloalkyl, (C_{6-10}) bicycloalkenyl, or Het, wherein said groups are unsubstituted or mono- or disubstituted by halogen, cyano, nitro, hydroxy, (C_{1-4}) alkyl and/or $O-(C_{1-4})$ alkyl, wherein the alkyl groups may be fluorinated.

16. The compound according to claim 15, wherein R^3 is cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl or cycloheptyl, or a group selected from



wherein all said groups are unsubstituted or substituted by fluorine, (C₁₋₃)alkyl or CF₃.

- 5 **17.** The compound according to claim 16, wherein **R³** is cyclopentyl or cyclohexyl.

- 10 **18.** The compound according to claim 1 wherein **R^{4a}**, **R^{4b}**, **R⁵** each are independently H, hydroxy, halogen, cyano, nitro, carboxyl, (C₁₋₄)alkyl, CF₃, (C₁₋₄)alkoxy, -O-(C₃₋₇)cycloalkyl, -O-(C₁₋₃)alkyl-(C₃₋₇)cycloalkyl, -O-aryl, -O-(C₁₋₃)alkyl-aryl, -O-Het, -O-(C₁₋₃)alkyl-Het, **NR^{N1}R^{N2}**, **COR^O**, **NR^{N2}COR^C**, **CONR^{N2}R^{N1}**, or **NR^{N3}CONR^{N1}R^{N2}**;
wherein **Het**, **R^C**, **R^O**, **R^{N1}**, **R^{N2}**, **R^{N3}** and **R¹⁶⁰** are as defined in claim 1; and wherein all said alkyl groups, including alkoxy, may be mono-, di- or trisubstituted by fluorine or mono-substituted by chlorine or bromine.
- 15 **19.** The compound according to claim 18 wherein **R^C**, **R^O** and **R^{N1}** are independently of each other H, (C₁₋₄)alkyl, aryl, (C₁₋₃)alkyl-aryl; wherein aryl is defined as phenyl optionally substituted with **R¹⁶⁰**, wherein **R¹⁶⁰** is defined as in claim 18; and
20 wherein all said alkyl groups may be mono-, di- or trisubstituted by fluorine or mono-substituted by chlorine or bromine; and
wherein **R^{N2}** and **R^{N3}** are independently H or methyl.

- 25 **20.** The compound according to claim 18 wherein **R^{4a}**, **R^{4b}**, **R⁵** each are independently H, hydroxy, halogen, cyano, nitro, methyl, CF₃, methoxy, carboxy, amino, -NMe₂, -CONH₂, -NHCONH₂, -CO-NHMe, -NHCONHMe, -CO-NMe₂ or -NHCONMe₂.

- 30 **21.** The compound according to claim 20 wherein **R^{4a}**, **R^{4b}**, **R⁵** each are H, methyl or methoxy.

- 22.** The compound according to claim 1 wherein **R^{4a}** is H or methyl.

23. The compound according to claim 1 wherein at least two of the substituents selected from R^{4a} , R^{4b} , R^5 are H.
24. The compound according to claim 1, wherein R^{60} is each defined as 1 to 4 substituents independently selected from:
- 5
- 1 to 3 substituents selected from halogen;
 - one of each substituent selected from: NO_2 , cyano, azido; and
 - 1 to 3 substituents selected from:
- 10
- a) (C_{1-4}) alkyl, (C_{3-7}) cycloalkyl, (C_{2-4}) alkenyl, (C_{2-4}) alkynyl, (C_{1-3}) alkyl- (C_{3-7}) cycloalkyl, all of which optionally being substituted with R^{150} ;
 - b) OR^O ;
 - e) $N(R^{N2})R^{N1}$;
 - f) $N(R^{N2})COR^C$;
 - j) $COOR^O$;
 - 15 k) $CON(R^{N2})R^{N1}$;
 - l) phenyl, Het, (C_{1-3}) alkyl)phenyl or (C_{1-3}) alkyl)Het; wherein Het is selected from furan, tetrahydrofuran, thiophene, tetrahydrothiophene, tetrahydropyran, pyridinyl, azetidine, pyrrolidine, piperidine, piperazine, morpholine, thiomorpholine, homopiperidine and
 - 20 homopiperazine, all of which optionally being substituted with R^{150} ;
- wherein said R^{N1} , R^C and/or R^O are optionally substituted with R^{150} as defined, and R^{150} , R^{N1} , R^{N2} , R^C and R^O are defined as in claim 1.
25. The compound according to claim 1, wherein
- 25 R^{150} is defined as 1 to 4 substituents independently selected from:
- 1 to 3 fluorine-substituents;
 - one of each substituent selected from: chlorine, bromine, iodine, NO_2 , cyano, azido; and
 - 1 to 3 substituents selected from:
- 30
- a) (C_{1-3}) alkyl, CF_3 , (C_{3-6}) cycloalkyl, (C_{1-3}) alkyl- (C_{3-6}) cycloalkyl, all of which optionally substituted with R^{160} ;
 - b) OR^O ;
 - e) $N(R^{N2})R^{N1}$;
 - f) $N(R^{N2})COR^C$;

j) COOR^{O} ;

k) $\text{CON}(\text{R}^{\text{N2}})\text{R}^{\text{N1}}$;

wherein said R^{N1} , R^{C} and/or R^{O} are optionally substituted with R^{160} as defined; and

5 R^{160} , R^{N1} , R^{N2} , R^{C} and R^{O} are defined as in claim 1.

26. The compound according to claim 1, wherein

R^{160} is defined as 1, 2 or 3 substituents independently selected from:

- 1, 2 or 3 fluorine substituents; and
- 10 - one of each substituent selected from chlorine, bromine, iodine, CN, nitro, methyl, trifluoromethyl, ethyl, n-propyl, i-propyl, COOH , COOCH_3 , OH , OCH_3 , OCF_3 , NH_2 , NHCH_3 , $\text{N}(\text{CH}_3)_2$, SO_2NH_2 , $\text{SO}_2\text{NHCOCH}_3$, NHCOCH_3 or CONH_2 , CONHCH_3 and $\text{CON}(\text{CH}_3)_2$.

15 27. The compound according to claim 1, wherein

R^{O} , R^{C} are independently defined as (C_{1-4}) alkyl, (C_{3-6}) cycloalkyl, (C_{1-3}) alkyl- (C_{3-6}) cycloalkyl, phenyl, benzyl, **Het**, (C_{1-3}) alkyl-**Het**; all of which are optionally substituted as defined; and R^{O} may also be H;

20

R^{N1} is H, (C_{1-4}) alkyl, (C_{3-6}) cycloalkyl, (C_{1-3}) alkyl- (C_{3-6}) cycloalkyl, phenyl, benzyl, phenylethyl, **Het**, (C_{1-3}) alkyl-**Het**; wherein said alkyl, cycloalkyl, alkyl-cycloalkyl, phenyl, benzyl, phenylethyl, **Het** and alkyl-**Het** are optionally substituted as defined; or

25

R^{N2} , R^{N3} , R^{N4} are independently H, methyl, ethyl, n-propyl, i-propyl, cyclopropyl, cyclopropylmethyl; all of which being optionally substituted with fluorine, carboxy or methoxycarbonyl; and/or wherein said ethyl, n-propyl or i-propyl is optionally substituted with hydroxy, methyl, methoxy, amino, $-\text{NH}(\text{CH}_3)$ and/or $-\text{N}(\text{CH}_3)_2$; and

30

in the case

a) of a group $\text{N}(\text{R}^{\text{N2}})\text{R}^{\text{N1}}$ the substituents R^{N2} and R^{N1} or

b) of a group $\text{NR}^{\text{N3}}-\text{N}(\text{R}^{\text{N2}})\text{R}^{\text{N1}}$ the substituents R^{N3} and R^{N1} or R^{N2} and R^{N1}

may be covalently bonded together to form a 5-, 6- or 7-membered saturated heterocycle which may have additionally one heteroatom selected from O, N, and S, wherein said heterocycle is optionally substituted as defined;

5 wherein **Het** is defined as in claim 1.

28. Use of a compound of the formula I according to claim 1, or a pharmaceutically acceptable salt thereof, as an inhibitor of HCV polymerase.

10 29. Use of a compound of the formula I according to claim 1, or a pharmaceutically acceptable salt thereof, as an inhibitor of RNA dependent RNA polymerase activity of the enzyme NS5B, encoded by HCV.

15 30. Use of a compound of the formula I according to claim 1, or a pharmaceutically acceptable salt thereof, as an inhibitor of HCV replication.

20 31. A method of treating or preventing HCV infection in a mammal, comprising administering to the mammal an effective amount of a compound of formula I according to claim 1, or a pharmaceutically acceptable salt thereof.

32. A method of treating or preventing HCV infection in a mammal, comprising administering to the mammal an effective amount of a compound of formula I according to claim 1, or a pharmaceutically acceptable salt thereof in combination with another antiviral agent.

25 33. A pharmaceutical composition for the treatment or prevention of HCV infection, comprising an effective amount of a compound of formula I according to claim 1, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.

30 34. The composition according to claim 33 further comprising a therapeutically effective amount of one or more antiviral agents.

35. The composition according to claim 34, wherein said antiviral agent is selected from: ribavirin and amantadine.
- 5 36. The composition according to claim 34 wherein the antiviral agent is an other anti-HCV agent.
- 10 37. The pharmaceutical composition according to claim 36, wherein the other anti-HCV agent is an immunomodulatory agent, in particular selected from β -, δ - γ -, and ω -interferon.
38. A composition according to claim 36, wherein said anti-HCV agent is another inhibitor of HCV polymerase.
- 15 39. The composition according to claim 36, wherein the other anti-HCV agent is an inhibitor of HCV NS3 protease.
40. The composition according to claim 36, wherein the other anti-HCV agent is an inhibitor of another target in the HCV life cycle.
- 20 41. A composition according to claim 40, wherein said inhibitor of another target in the HCV life cycle is an agent that inhibits a target selected from HCV helicase, HCV NS2/3 protease and HCV IRES.
- 25 42. Use of a compound of formula I according to claim 1, or of a pharmaceutically acceptable salt thereof, for the manufacture of a medicament for the treatment and/or the prevention of a viral infection, preferably an HCV infection.